

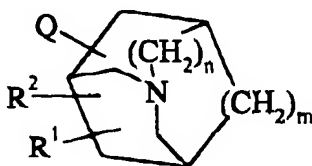
## AMENDMENTS

## Amendments to the claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

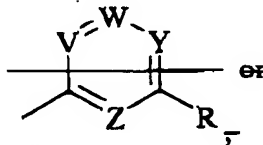
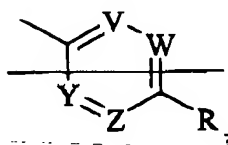
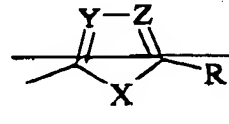
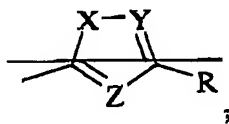
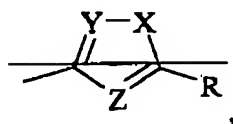
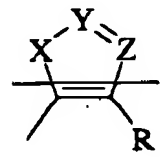
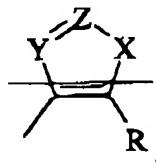
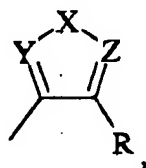
## Listing of claims:

1. (currently amended) A pharmaceutical composition comprising at least one M4 selective muscarinic agonist selected from the azacyclic ring system having the formula I

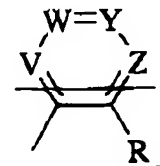


I

including and geometrical isomers, enantiomers, diastereomers, racemates, acid addition salts, salts thereof with a pharmaceutically acceptable acid, and prodrugs thereof, wherein Q is



or



X is ~~CH<sub>2</sub>, NH, O or S~~;

V, W, Y and Z ~~independently~~ are CH or N;

n and m independently are 0, 1, 2, 3 or 4;

$R^1$  and  $R^2$  are at any position on the azacyclic ring, including the point of attachment of the heterocycle Q, and independently are hydrogen, -OH, halogen, -NH<sub>2</sub>, carboxy, straight or branched C<sub>1-10</sub>-alkyl, C<sub>1-10</sub>-alkenyl, or C<sub>1-10</sub>-alkynyl, straight or branched C<sub>1-10</sub>-alkoxy, or straight or branched C<sub>1-10</sub>-alkyl substituted with -OH, -CN, -CHO, -OH, -OR<sup>3</sup>, -SR<sup>3</sup>, -NH<sub>2</sub>, -NHR<sup>3</sup>, -NR<sup>3</sup>R<sup>4</sup>, -NO<sub>2</sub>, -SOR<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, -COR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONH<sub>2</sub>, -CONHR<sup>3</sup>, -CONR<sup>3</sup>R<sup>4</sup>, or -CH=NOR<sup>3</sup>; or

$R^1$  and  $R^2$  independently are phenyl, phenoxy, benzoyl, benzyl or benzyloxycarbonyl, each of which are unsubstituted or substituted with halogen, -CN, C<sub>1-10</sub>-alkyl, C<sub>1-10</sub>-alkoxy, or C<sub>1-10</sub>-alkylthio;

R is hydrogen, halogen, -CN, -CHO, -OH, -OR<sup>3</sup>, -SR<sup>3</sup>, -NH<sub>2</sub>, -NHR<sup>3</sup>, -NR<sup>3</sup>R<sup>4</sup>, -NO<sub>2</sub>, -SOR<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, -COR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONH<sub>2</sub>, -CONHR<sup>3</sup>, -CONR<sup>3</sup>R<sup>4</sup>, or -CH=NOR<sup>3</sup>; or

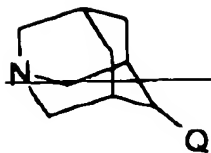
R is phenyl, phenoxy, benzoyl, benzyl or benzyloxycarbonyl, each of which are unsubstituted or substituted with halogen, -CN, C<sub>1-15</sub>-alkyl, C<sub>1-10</sub>-alkoxy, or C<sub>1-10</sub>-alkylthio; or

R is a 5 or 6 membered saturated, partly saturated or aromatic heterocyclic ring containing one to three heteroatoms; and

$R^3$  and  $R^4$  independently are straight, branched, or cyclic C<sub>1-15</sub>-alkyl, C<sub>2-15</sub>-alkenyl, C<sub>2-15</sub>-alkynyl, or combinations thereof, or  $R^3$  and  $R^4$  independently are phenyl, phenoxy, benzoyl, benzyl or benzyloxycarbonyl groups, each of which are unsubstituted or substituted with H, halogen, -CN, C<sub>1-15</sub>-alkyl, C<sub>1-10</sub>-alkoxy, C<sub>1-10</sub>-alkylthio, or aryl; or

$R^3$  and  $R^4$  independently are 5 or 6 membered saturated, partly saturated or aromatic heterocyclic rings containing one to three heteroatoms; and further comprising one or more additional analgesics.

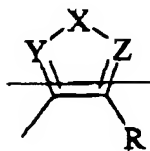
2. (currently amended) The composition according to claim 1 wherein in formula I of the M4 selective muscarinic agonist n and m both are 1 and the azacyclic ring system has the structural formula:



H

wherein

Q is:



X is S,

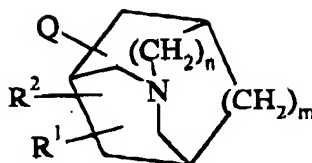
Y and Z are N, and

R is OR<sup>3</sup> or SR<sup>3</sup>.

3. (original) The composition according to claim 2 wherein R<sup>3</sup> of the M4 selective muscarinic agonist is -CH<sub>3</sub>, -CH<sub>2</sub>CH<sub>3</sub>, -CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub> or -CH<sub>2</sub>CH(CH<sub>3</sub>)<sub>2</sub>.
4. (original) The composition according to claim 1 wherein the M4 selective muscarinic agonist is selected from the group consisting of
- 3-(5-Aza-2-chlorotricyclo[3.3.1.1<3,7>]dec-2-yl)-4-chloro-1,2,5-thiadiazole;
  - 3-(5-Azatricyclo[3.3.1.1<3,7>]dec-2-yl)-4-chloro-1,2,5-thiadiazole;
  - 3-(5-azatricyclo[3.3.1.1<3,7>]dec-2-yl)-4-methoxy-1,2,5-thiadiazole;
  - 3-(5-azatricyclo[3.3.1.1<3,7>]dec-2-yl)-4-ethoxy-1,2,5-thiadiazole;
  - 3-(5-azatricyclo[3.3.1.1<3,7>]dec-2-yl)-4-propoxy-1,2,5-thiadiazole;
  - 3-(5-azatricyclo[3.3.1.1<3,7>]dec-2-yl)-4-butoxy-1,2,5-thiadiazole;
  - 3-(5-azatricyclo[3.3.1.1<3,7>]dec-2-yl)-4-(cyclopropylmethoxy)-1,2,5-thiadiazole; and
  - 3-(5-azatricyclo[3.3.1.1<3,7>]dec-2-yl)-4-(2-methyl-propoxy)-1,2,5-thiadiazole;
  - 4-[4-(propylsulfanyl)-1,2,5-thiadiazol-3-yl]-1-azatricyclo[3.3.1.1<3,7>]decane hydrochloride
  - 4-[4-(methylsulfanyl)-1,2,5-thiadiazol-3-yl]-1-azatricyclo[3.3.1.1<3,7>]decane
  - 4-[4-(ethylsulfanyl)-1,2,5-thiadiazol-3-yl]-1-azatricyclo[3.3.1.1<3,7>]decane

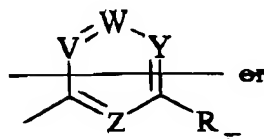
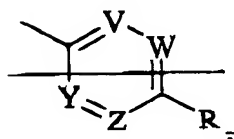
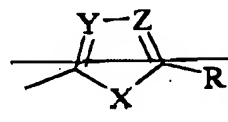
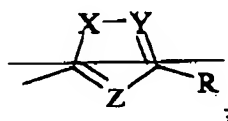
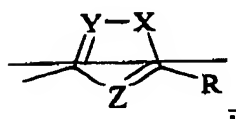
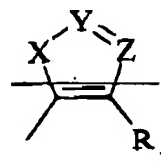
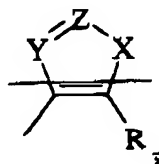
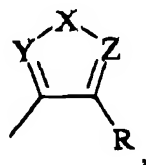
- l) 4-[4-(butylsulfanyl)-1,2,5-thiadiazol-3-yl]-1-azatricyclo[3.3.1.1<3,7>]decane  
m) 4-[4-(2-methyl-propylsulfanyl)-1,2,5-thiadiazol-3-yl]-1-azatricyclo[3.3.1.1<3,7>]decane  
n) 4-[4-(cyclopropylmethylsulfanyl)-1,2,5-thiadiazol-3-yl]-1-azatricyclo[3.3.1.1<3,7>]decane.

5. (original) The composition according to claim 4 wherein the M4 selective muscarinic agonist is 4-s-[4-(propylsulfanyl)-1,2,5-thiadiazol-3-yl]-1-azatricyclo[3.3.1.1<3,7>]decane hydrochloride.
6. (original) The composition according to claim 1 further comprising a pharmaceutically acceptable carrier.
7. (original) The composition according to claim 1 wherein the additional analgesic is selected from the group of opioid analgesics, nonsteroidal anti-inflammatory drugs and other analgesics.
8. (original) The composition according to claim 7 wherein the additional analgesic is an opioid analgesic.
9. (original) The composition according to claim 8 wherein the opioid analgesic is selected from the group of morphine and codeine.
10. (original) The composition according to claim 7 wherein the additional analgesic is a non-steroidal anti-inflammatory drug.
11. (original) The composition according to claim 10 wherein the non-steroidal anti-inflammatory drug is selected from the group of acetaminophen, ibuprofen, celecoxib and rofecoxib.
12. (original) The composition according to claim 7 wherein the additional analgesic is selected from the group of nicotinic agonists, NMDA antagonists, epileptics and alpha adrenoceptor agonists.
13. (withdrawn) A method of inducing analgesia, the method comprising co-administration of at least one M4 selective muscarinic agonist selected from the azacyclic ring system having the formula I

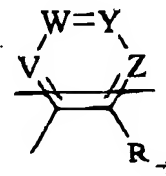


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including ~~and~~ geometrical isomers, enantiomers, diastereomers, racemates, acid addition salts, salts thereof with a pharmaceutically acceptable acid, and prodrugs thereof, wherein Q is



or



X is ~~CH<sub>2</sub>, NH, O or S~~;

V, W, Y and Z ~~independently~~ are CH or N;

n and m ~~independently~~ are 0, 1, 2, 3 or 4;

R<sup>1</sup> and R<sup>2</sup> are at any position on the azacyclic ring, including the point of attachment of the heterocycle Q, and independently are hydrogen, -OH, halogen, -NH<sub>2</sub>, carboxy, straight or branched C<sub>1-10</sub>-alkyl, C<sub>1-10</sub>-alkenyl, or C<sub>1-10</sub>-alkynyl, straight or branched C<sub>1-10</sub>-alkoxy, or straight or branched C<sub>1-10</sub>-alkyl substituted with -OH, -CN, -CHO, -OH, -OR<sup>3</sup>, -SR<sup>3</sup>, -NH<sub>2</sub>, -NHR<sup>3</sup>, -NR<sup>3</sup>R<sup>4</sup>, -NO<sub>2</sub>, -SOR<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, -COR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONH<sub>2</sub>, -CONHR<sup>3</sup>, -CONR<sup>3</sup>R<sup>4</sup>, or -CH=NOR<sup>3</sup>; or

$R^1$  and  $R^2$  independently are phenyl, phenoxy, benzoyl, benzyl or benzyloxycarbonyl, each of which are unsubstituted or substituted with halogen, -CN,  $C_{1-10}$ -alkyl,  $C_{1-10}$ -alkoxy, or  $C_{1-10}$ -alkylthio;

R is hydrogen, halogen, -CN, -CHO, -OH, -OR<sup>3</sup>, -SR<sup>3</sup>, -NH<sub>2</sub>, -NHR<sup>3</sup>, -NR<sup>3</sup>R<sup>4</sup>, -NO<sub>2</sub>, -SOR<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, -COR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONH<sub>2</sub>, -CONHR<sup>3</sup>, -CONR<sup>3</sup>R<sup>4</sup>, or -CH=NOR<sup>3</sup>; or

R is phenyl, phenoxy, benzoyl, benzyl or benzyloxycarbonyl, each of which are unsubstituted or substituted with halogen, -CN,  $C_{1-15}$ -alkyl,  $C_{1-10}$ -alkoxy, or  $C_{1-10}$ -alkylthio; or

R is a 5 or 6 membered saturated, partly saturated or aromatic heterocyclic ring containing one to three heteroatoms; and

$R^3$  and  $R^4$  independently are straight, branched, or cyclic  $C_{1-15}$ -alkyl,  $C_{2-15}$ -alkenyl,  $C_{2-15}$ -alkynyl, or combinations thereof, or  $R^3$  and  $R^4$  independently are phenyl, phenoxy, benzoyl, benzyl or benzyloxycarbonyl groups, each of which are unsubstituted or substituted with H, halogen, -CN,  $C_{1-15}$ -alkyl,  $C_{1-10}$ -alkoxy,  $C_{1-10}$ -alkylthio, or aryl; or

$R^3$  and  $R^4$  independently are 5 or 6 membered saturated, partly saturated or aromatic heterocyclic rings containing one to three heteroatoms; with one or more additional analgesics.

14. (withdrawn) A method of inducing analgesia according to claim 13, the method comprising administering an analgesia-inducing amount of a composition according to claim 1 to a mammal in need thereof.
15. (canceled) A composition according to claim 1 for use as a medicament.
16. (canceled) A composition according to claim 1 for use as an analgesic.
17. (canceled) Use of the composition according to claim 1 for the manufacture of a medicament for treatment of analgesia.